	Application No.	Applicant(s)
	09/718,717	JAKOBOVITS ET AL.
Notice of Allowability	Examiner	Art Unit
	Terry A. McKelvey	1636
The MAILING DATE of this communication appe All claims being allowable, PROSECUTION ON THE MERITS IS (herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIG	(OR REMAINS) CLOSED in or other appropriate commu GHTS. This application is s	this application. If not included
1. This communication is responsive to the reply filed 10/22/0	<u>4</u> .	
2. The allowed claim(s) is/are 11 and 12.		
3. \boxtimes The drawings filed on <u>22 November 2000</u> are accepted by the second sec	the Examiner.	
 4. ☐ Acknowledgment is made of a claim for foreign priority unda) ☐ All b) ☐ Some* c) ☐ None of the: 1. ☐ Certified copies of the priority documents have 2. ☐ Certified copies of the priority documents have 3. ☐ Copies of the certified copies of the priority documents have International Bureau (PCT Rule 17.2(a)). * Certified copies not received: 	been received. been received in Applicatio	n No
Applicant has THREE MONTHS FROM THE "MAILING DATE" of	of this communication to file	a souly according with the convince
noted below. Failure to timely comply will result in ABANDONMI THIS THREE-MONTH PERIOD IS NOT EXTENDABLE.	ENT of this application.	a reply complying with the requirements
5. A SUBSTITUTE OATH OR DECLARATION must be submit INFORMAL PATENT APPLICATION (PTO-152) which give:	tted. Note the attached EXA s reason(s) why the oath or	MINER'S AMENDMENT or NOTICE OF declaration is deficient.
6. CORRECTED DRAWINGS (as "replacement sheets") must	be submitted.	
(a) 🔲 including changes required by the Notice of Draftsperson's Patent Drawing Review (PTO-948) attached		
1) 🗌 hereto or 2) 🔲 to Paper No./Mail Date		
(b) including changes required by the attached Examiner's Paper No./Mail Date	Amendment / Comment or	in the Office action of
Identifying indicia such as the application number (see 37 CFR 1.8 each sheet. Replacement sheet(s) should be labeled as such in the	84(c)) should be written on the header according to 37 CFI	e drawings in the front (not the back) of R 1.121(d).
7. DEPOSIT OF and/or INFORMATION about the depos attached Examiner's comment regarding REQUIREMENT F	it of BIOLOGICAL MATE OR THE DEPOSIT OF BIO	RIAL must be submitted. Note the LOGICAL MATERIAL.
•		
Attachment(s)		
1. Notice of References Cited (PTO-892)	5. Notice of Inf	ormal Patent Application (PTO-152)
2. Notice of Draftperson's Patent Drawing Review (PTO-948)		mmary (PTO-413),
3. Information Disclosure Statements (PTO-1449 or PTO/SB/08 Paper No./Mail Date		Mail Date Amendment/Comment
4. Examiner's Comment Regarding Requirement for Deposit	8. 🗌 Examiner's S	Statement of Reasons for Allowance
of Biological Material	9. 🗌 Other	

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EXAMINER'S AMENDMENT

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An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with Paula Borden on 12/14/04.

The application has been amended as follows:

In the title:

The title has been replaced with the following:

-- Method for preparing a mammalian cell deficient in HPRT

In the specification:

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At page 3, the three paragraphs from line 3 to line 18 have been replaced with the following:

-- FIG. 1 depicts the hprt locus in normal mouse and mutant E14TG2a cells and targeting constructs used to delete 55 kb, as described in Example 1, infra (Restriction enzyme sites: B; BamHI, C; ClaI, E; EcoRI, X; XhoI, SS; SaII and XmnI).

FIGS. 2A-2B and 3A to 3D depict pulse field gel electrophoresis (PFGE) of hprt gene locus in E14.1, E14TG2a, pm4.2 and pm5x.1 cells, as described in Example 1, infra. FIGS. 2A and 2B provide schematic illustrations of SacII and BssHII restriction sites in the hprt gene locus in E14.1 and pm4.2ES cells. In FIGS. 3A to 3D, a=E14.1; b=E14TG2a; c=pm4.2 and d=pm5x.1 in each panel DNA was digested with SacII (FIGS. 3A, 3B) or BssHII (FIGS. 3C, 3D) and probed with 5' probe (FIGS. 3A or 3C) or 3' probe (FIGS. 3B or 3D).

FIG. 4 depicts inactivation of these mouse hprt locus by gene-targeted 55 kb deletion, using the targeting construct (pMp4), as described in Example 1, infra. The probes used for analysis (5' and 3' probe) and the structure of the hprt gene around targeted region in E14.1 cells, E14TG2a cells and

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targeted clones obtained: B; BamHI, C; ClaI, X; XhoI, R; RsaI, jx; junction and Ex; exon.

FIGS. 5A to 5D provide the results of Southern blot analysis of hprt gene locus in E14.1 (column a) E14TG2a (column b); and targeted ES clones pm4.2 (column c) and pm5x.1 (column d). DNA was digested with BamHI (FIGS. 5A and 5B) or with BamHI/C1aI (FIGS. 5C and 5D) and was probed with 5' probe (FIGS. 5A and 5C) or with 3' probe (FIGS. 5B and 5D). --

In the claims:

- 12. (Currently amended) The method of claim 11 which further comprises the steps of
- a) identifying cells containing said deletion deficiency by selecting cells containing a selectable marker present in said construct; and
- b) recovering cells containing said deletion deficiency.

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Conclusion

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to (571) 272-0547.

Patent applicants with problems or questions regarding electronic images that can be viewed in the Patent Application Information Retrieval system (PAIR) can now contact the USPTO's Patent Electronic Business Center (Patent EBC) for assistance. Representatives are available to answer your questions daily from 6 am to midnight (EST). The toll free number is (866) 217-9197. When calling please have your application serial or patent number, the type of document you are having an image problem with, the number of pages and the specific nature of the problem. The Patent Electronic Business Center will notify applicants of the resolution of the problem within 5-7 business days. Applicants can also check PAIR to confirm that the problem has been corrected. The USPTO's Patent Electronic Business Center is a complete service center supporting all patent business on the Internet. The USPTO's PAIR system provides Internet-based access to patent application status and history information. It also enables applicants to view the scanned images of their own application file folder(s) as well as general patent information available to the public.

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Any inquiry concerning rejections or objections in this communication or earlier communications from the examiner should be directed to Terry A. McKelvey whose telephone number is (571) 272-0775. The examiner can normally be reached on Monday through Friday, except for Wednesdays, from about 7:30 AM to about 6:00 PM. A phone message left at this number will be responded to as soon as possible (i.e., shortly after the examiner returns to his office).

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Dr. Remy Yucel can be reached at (571) 272-0781.

Jenga Mc Keller

Terry A. McKelvey, Ph.D. Primary Examiner Art Unit 1636

December 27, 2004